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## **Clinical Evaluation of Streptococcus pneumoniae Polymerase Chain Reaction in Children with Suspected Septicemia**

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# The Pediatric Infectious Disease Journal

## Clinical evaluation of Streptococcus pneumoniae PCR in children with suspected septicaemia --Manuscript Draft--

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Corresponding Author:	Thomas Walter Bourke, M.D. Royal Belfast Hospital for Sick Children Belfast, UNITED KINGDOM
Corresponding Author Secondary Information:	
Corresponding Author's Institution:	Royal Belfast Hospital for Sick Children
Corresponding Author's Secondary Institution:	
First Author:	Thomas Walter Bourke, M.D.
First Author Secondary Information:	
Order of Authors:	Thomas Walter Bourke, M.D.
	Derek James Fairley, PhD
	James Patrick McKenna, PhD
	Peter Valentine Coyle, MD
	Michael David Shields, MD
Order of Authors Secondary Information:	
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**Clinical evaluation of *Streptococcus pneumoniae* PCR in children with suspected septicaemia.**

Thomas Walter Bourke MD <sup>a,c</sup>

Derek James. Fairley PhD <sup>a,b</sup>

James Patrick McKenna PhD <sup>b</sup>

Prof Peter Valentine Coyle MD <sup>b</sup>

Prof Michael David Shields MD <sup>a,c</sup>

<sup>a</sup> Centre for Infection and Immunity, Queen's University Belfast, Belfast, UK

<sup>b</sup> Regional Virus Laboratory, Department of Microbiology, Belfast Health & Social Care Trust, Belfast, UK

<sup>c</sup> Royal Belfast Hospital for Sick Children, Belfast Health & Social Care Trust, Belfast, UK

Address correspondence to:

Dr Thomas Walter Bourke

Royal Belfast Hospital for Sick Children, Falls Road, Belfast, United Kingdom. BT12 6BE.

Telephone 0044 28 90634094. Fax 0044 28 90632816.

tombourke@doctors.net.uk

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Manuscript

Invasive infection with *Streptococcus pneumoniae* is the second commonest cause of meningitis and septicaemia in childhood, resulting in 10 to 15% of cases in the UK<sup>1,2</sup>. Isolation of *Streptococcus pneumoniae* from a normally sterile site remains the gold standard for diagnosis. Blood culture sensitivity is low (around 45%) and decreases further when antibiotics have been administered (around 20%)<sup>3</sup>. Despite its accuracy and potential to detect pneumococcus in culture negative patients, the use of molecular testing for *Streptococcus pneumoniae* is not widespread and not yet recommended as part of routine investigations of suspected septicaemia<sup>1,2</sup>.

We retrospectively tested 117 blood specimens for *Streptococcus pneumoniae lytA* using previously published PCR assays<sup>4</sup>. These specimens were collected from children with suspected meningitis or septicaemia as part of an ethically approved study on the diagnostic accuracy of *Neisseria meningitidis* Loop Mediated Isothermal Assay<sup>5</sup>.

Four of the 117 children tested had pneumococcal *lytA* detected by PCR. Only two of these had invasive pneumococcal disease confirmed by blood culture. The two children who were culture negative had clinical signs of septicaemia: A 10 month old boy with fever vomiting and listlessness who required fluid resuscitation and admission to intensive care; A two year old girl with fever, cough, poor feeding, poor perfusion and focal chest signs.

Evaluating *lytA* PCR as a confirmatory test for pneumococcus is complicated because it appears to perform better than the gold standard traditional culture. The children included in this study were defined as 'those whom the attending clinician suspected as having meningitis or septicaemia'. Review of the clinical details of the children in this study in combination with a positive *lytA* PCR make it likely that these children had invasive pneumococcal disease which conventional culture methods failed to

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diagnose. Although our study was small and pneumococcal disease was rare these data suggest that addition of *lytA* PCR to the routine investigations of children presenting with signs of septicaemia and meningitis is likely to greatly improve laboratory diagnosis of this serious infection. This is clinically a very desirable outcome as treatment for invasive pneumococcal disease requires a longer course of treatment than meningococcal disease<sup>2</sup>. We believe that reviewed National Institute for Health and Care Excellence and other guidelines should consider recommending this test to the panel of investigations of children with suspected meningitis or septicaemia.

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